

**To:** Christine Olinger[Olinger.Christine@epa.gov]; Donna Davis[Davis.Donna@epa.gov]; Swartz, Christina[Swartz.Christina@epa.gov]; Dana Vogel[Vogel.Dana@epa.gov]; Jeffrey Dawson (Dawson.Jeff@epa.gov)[Dawson.Jeff@epa.gov]  
**Cc:** Wade Britton (Britton.Wade@epa.gov)[Britton.Wade@epa.gov]  
**From:** Lowit, Anna  
**Sent:** Wed 11/2/2016 6:51:57 PM  
**Subject:** FW: Carbaryl call with Scitovation today  
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Here is the information that Cecilia needs for developing scenarios using pbpk models, see below.

Wade & Cecilia have done this for cpfos, so this is the best place to start.---pp 62-65 in this file.

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**From:** Tan, Cecilia  
**Sent:** Wednesday, November 02, 2016 1:38 PM  
**To:** Lowit, Anna <Lowit.Anna@epa.gov>  
**Cc:** Walsh, Linsey <Walsh.Linsey@epa.gov>; Gallagher, Sarah <Gallagher.Sarah@epa.gov>; Godshall, Joshua <Godshall.Joshua@epa.gov>; Olinger, Christine <Olinger.Christine@epa.gov>  
**Subject:** RE: Carbaryl call with Scitovation today

Thank you, Anna, for your quick responses.

The only information they need for the list of scenarios include the following:

## Ex. 5 - Deliberative Process

Here is one example: 4.8 kg, single daily oral dose for 21 days.

They will be able to finish the oral PoD sooner than other scenarios (e.g., worker).

Cecilia

**From:** Lowit, Anna

**Sent:** Wednesday, November 02, 2016 1:11 PM

**To:** Tan, Cecilia <[Tan.Cecilia@epa.gov](mailto:Tan.Cecilia@epa.gov)>

**Cc:** Walsh, Linsey <[Walsh.Linsey@epa.gov](mailto:Walsh.Linsey@epa.gov)>; Gallagher, Sarah <[Gallagher.Sarah@epa.gov](mailto:Gallagher.Sarah@epa.gov)>;  
Godshall, Joshua <[Godshall.Joshua@epa.gov](mailto:Godshall.Joshua@epa.gov)>; Olinger, Christine

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**Subject:** Re: Carbaryl call with Scitovation today

Cecilia

Remember from our meeting yesterday, in prep for the sap discussion meeting with registrants and developers we (with help from wade and the teams) will set up a template of pods that we

will expect them to do---you won't be doing it. But, yes, 10% ache inhibition.

One of the BCs is going to help get the list of scenarios going. I've cc'd josh here who knows carbaryl.

Sent from my iPhone

On Nov 2, 2016, at 12:27 PM, Tan, Cecilia <[Tan.Cecilia@epa.gov](mailto:Tan.Cecilia@epa.gov)> wrote:

Hi, Anna,

Here is a short summary of our conference call with ScitoVation today.

They are updating the model to include life-stage specific parameters, and the oral model should be ready before the end of December (they also said it could be done within a month). They will add the dermal and inhalation routes, and those will be ready in January.

I think their approach for including life-stage changes is reasonable. Not sure if there are existing dermal or inhalation data for them to evaluate the model, but we will wait and see.

Here are three questions for you and the team:

1. I am guessing that we will use the carbaryl model to estimate PoDs based on 10% RBC AChE inhibition (and thus short-term exposure)? Are we also interested in using the model for cancer endpoint?
2. I asked them about their plan for peer review. I made it clear that we are not requiring one, but I just want to know their plan. I will definitely examine the model structure, code, and parameter values. An additional peer review will save me a lot of time and give this model more credibility. They will discuss the plan for the review (yes/no, timeline) and let us know. My question for you is: will additional peer review be helpful?
3. If we are going to estimate PoDs for different scenarios based on AChE endpoint, will Wade help putting together the list? I don't think that they need those scenarios before the model is finalized, even though they keep asking for them. I plan to ask them to run a single daily oral exposure for 30 days for an adult to see how long it will take for the model to run. My experience is that it will take probably 5 – 6 runs to find the PoD based on AChE, so we can multiply the time to run one scenario by 6 by the number of scenario to get an estimate of total modeling time.

Lindsey, Sarah, and Josh (sorry, I forgot Josh's last name so I didn't cc him) were also on the phone. If I missed anything, please feel free to add.

Thank you.

Cecilia